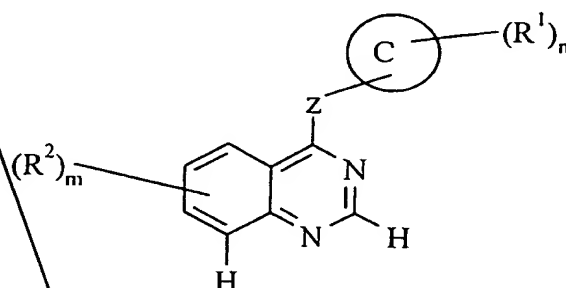


Claims:

1. The use of a compound of the formula I:



(I)

wherein:

ring C is an 8, 9, 10, 12 or 13-membered bicyclic or tricyclic moiety which moiety may be saturated or unsaturated, which may be aromatic or non-aromatic, and which optionally may contain 1-3 heteroatoms selected independently from O, N and S;

Z is -O-, -NH-, -S-, -CH₂- or a direct bond;

n is an integer from 0 to 5;

m is an integer from 0 to 3;

R² represents hydrogen, hydroxy, halogeno, cyano, nitro, trifluoromethyl, C₁₋₃alkyl, C₁₋₃alkoxy, C₁₋₃alkylsulphanyl, -NR³R⁴ (wherein R³ and R⁴, which may be the same or different, each represents hydrogen or C₁₋₃alkyl), or R⁵X¹- (wherein X¹ represents a direct bond, -O-, -CH₂-, -OC(O)-, -C(O)-, -S-, -SO-, -SO₂-, -NR⁶C(O)-, -C(O)NR⁷-, -SO₂NR⁸-, -NR⁹SO₂- or -NR¹⁰- (wherein R⁶, R⁷, R⁸, R⁹ and R¹⁰ each independently represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl), and R⁵ is selected from one of the following twenty-two groups:

1) hydrogen, oxiranylC₁₋₄alkyl or C₁₋₅alkyl which may be unsubstituted or which may be substituted with one or more groups selected from hydroxy, fluoro, chloro, bromo and amino;

2) C₁₋₅alkylX²C(O)R¹¹ (wherein X² represents -O- or -NR¹²- (in which R¹² represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl) and R¹¹ represents C₁₋₃alkyl, -NR¹³R¹⁴ or -OR¹⁵

(wherein R¹³, R¹⁴ and R¹⁵ which may be the same or different each represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl));

- 3) $C_{1,3}\text{alkyl}X^3R^{16}$ (wherein X^3 represents -O-, -S-, -SO-, -SO₂-, -OC(O)-, -NR¹⁷C(O)-, -C(O)NR¹⁸-, -SO₂NR¹⁹-, -NR²⁰SO₂- or -NR²¹- (wherein R¹⁷, R¹⁸, R¹⁹, R²⁰ and R²¹ each independently represents hydrogen, C_{1,3}alkyl or C_{1,3}alkoxyC_{2,3}alkyl) and R¹⁶ represents hydrogen, C_{1,3}alkyl, cyclopentyl, cyclohexyl or a 5-6-membered saturated heterocyclic group with 1-2 heteroatoms, selected independently from O, S and N, which C_{1,3}alkyl group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno and C_{1,4}alkoxy and which cyclic group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno, cyano, C_{1,4}cyanoalkyl, C_{1,4}alkyl, C_{1,4}hydroxyalkyl, C_{1,4}alkoxy, C_{1,4}alkoxyC_{1,4}alkyl, C_{1,4}alkylsulphonylC_{1,4}alkyl, C_{1,4}alkoxycarbonyl, C_{1,4}aminoalkyl, C_{1,4}alkylamino, di(C_{1,4}alkyl)amino, C_{1,4}alkylaminoC_{1,4}alkyl, di(C_{1,4}alkyl)aminoC_{1,4}alkyl, C_{1,4}alkylaminoC_{1,4}alkoxy, di(C_{1,4}alkyl)aminoC_{1,4}alkoxy and a group $-(O-)_f(C_{1,4}\text{alkyl})_g\text{ringD}$ (wherein f is 0 or 1, g is 0 or 1 and ring D is a 5-6-membered saturated heterocyclic group with 1-2 heteroatoms, selected independently from O, S and N, which cyclic group may bear one or more substituents selected from C_{1,4}alkyl));
- 4) $C_{1,3}\text{alkyl}X^4C_{1,3}\text{alkyl}X^5R^{22}$ (wherein X^4 and X^5 which may be the same or different are each -O-, -S-, -SO-, -SO₂-, -NR²³C(O)-, -C(O)NR²⁴-, -SO₂NR²⁵-, -NR²⁶SO₂- or -NR²⁷- (wherein R²³, R²⁴, R²⁵, R²⁶ and R²⁷ each independently represents hydrogen, C_{1,3}alkyl or C_{1,3}alkoxyC_{2,3}alkyl) and R²² represents hydrogen, C_{1,3}alkyl or C_{1,3}alkoxyC_{2,3}alkyl);
- 5) R²⁸ (wherein R²⁸ is a 5-6-membered saturated heterocyclic group (linked via carbon or nitrogen) with 1-2 heteroatoms, selected independently from O, S and N, which heterocyclic group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno, cyano, C_{1,4}cyanoalkyl, C_{1,4}alkyl, C_{1,4}hydroxyalkyl, C_{1,4}alkoxy, C_{1,4}alkoxyC_{1,4}alkyl, C_{1,4}alkylsulphonylC_{1,4}alkyl, C_{1,4}alkoxycarbonyl, C_{1,4}aminoalkyl, C_{1,4}alkylamino, di(C_{1,4}alkyl)amino, C_{1,4}alkylaminoC_{1,4}alkyl, di(C_{1,4}alkyl)aminoC_{1,4}alkyl, C_{1,4}alkylaminoC_{1,4}alkoxy, di(C_{1,4}alkyl)aminoC_{1,4}alkoxy and a group $-(O-)_f(C_{1,4}\text{alkyl})_g\text{ringD}$ (wherein f is 0 or 1, g is 0 or 1 and ring D is a 5-6-membered saturated heterocyclic group with 1-2 heteroatoms, selected independently from O, S and N, which cyclic group may bear one or more substituents selected from C_{1,4}alkyl));
- 6) C_{1,3}alkylR²⁸ (wherein R²⁸ is as defined herein);
- 7) C_{2,3}alkenylR²⁸ (wherein R²⁸ is as defined herein);
- 8) C_{2,3}alkynylR²⁸ (wherein R²⁸ is as defined herein);

- 317 -

- 9) R^{29} (wherein R^{29} represents a pyridone group, a phenyl group or a 5-6-membered aromatic heterocyclic group (linked via carbon or nitrogen) with 1-3 heteroatoms selected from O, N and S, which pyridone, phenyl or aromatic heterocyclic group may carry up to 5 substituents selected from hydroxy, halogeno, amino, C_{1-4} alkyl, C_{1-4} alkoxy, C_{1-4} hydroxyalkyl, C_{1-4} aminoalkyl, C_{1-4} alkylamino, C_{1-4} hydroxyalkoxy, carboxy, trifluoromethyl, cyano, - $C(O)NR^{30}R^{31}$, $-NR^{32}C(O)R^{33}$ (wherein R^{30} , R^{31} , R^{32} and R^{33} , which may be the same or different, each represents hydrogen, C_{1-4} alkyl or C_{1-3} alkoxy C_{2-3} alkyl) and a group $-(O)_f(C_{1-4}alkyl)_gringD$ (wherein f is 0 or 1, g is 0 or 1 and ring D is a 5-6-membered saturated heterocyclic group with 1-2 heteroatoms, selected independently from O, S and N, which cyclic group may bear one or more substituents selected from C_{1-4} alkyl));
- 10) $C_{1-3}alkylR^{29}$ (wherein R^{29} is as defined herein);
- 11) $C_{2-3}alkenylR^{29}$ (wherein R^{29} is as defined herein);
- 12) $C_{2-3}alkynylR^{29}$ (wherein R^{29} is as defined herein);
- 13) $C_{1-3}alkylX^6R^{29}$ (wherein X^6 represents $-O-$, $-S-$, $-SO-$, $-SO_2-$, $-NR^{34}C(O)-$, $-C(O)NR^{35}-$, $-SO_2NR^{36}-$, $-NR^{37}SO_2-$ or $-NR^{38}-$ (wherein R^{34} , R^{35} , R^{36} , R^{37} and R^{38} each independently represents hydrogen, $C_{1-3}alkyl$ or $C_{1-3}alkoxyC_{2-3}alkyl$) and R^{29} is as defined herein);
- 14) $C_{2-3}alkenylX^7R^{29}$ (wherein X^7 represents $-O-$, $-S-$, $-SO-$, $-SO_2-$, $-NR^{39}C(O)-$, $-C(O)NR^{40}-$, $-SO_2NR^{41}-$, $-NR^{42}SO_2-$ or $-NR^{43}-$ (wherein R^{39} , R^{40} , R^{41} , R^{42} and R^{43} each independently represents hydrogen, $C_{1-3}alkyl$ or $C_{1-3}alkoxyC_{2-3}alkyl$) and R^{29} is as defined herein);
- 15) $C_{2-3}alkynylX^8R^{29}$ (wherein X^8 represents $-O-$, $-S-$, $-SO-$, $-SO_2-$, $-NR^{44}C(O)-$, $-C(O)NR^{45}-$, $-SO_2NR^{46}-$, $-NR^{47}SO_2-$ or $-NR^{48}-$ (wherein R^{44} , R^{45} , R^{46} , R^{47} and R^{48} each independently represents hydrogen, $C_{1-3}alkyl$ or $C_{1-3}alkoxyC_{2-3}alkyl$) and R^{29} is as defined herein);
- 16) $C_{1-4}alkylX^9C_{1-4}alkylR^{29}$ (wherein X^9 represents $-O-$, $-S-$, $-SO-$, $-SO_2-$, $-NR^{49}C(O)-$, $-C(O)NR^{50}-$, $-SO_2NR^{51}-$, $-NR^{52}SO_2-$ or $-NR^{53}-$ (wherein R^{49} , R^{50} , R^{51} , R^{52} and R^{53} each independently represents hydrogen, $C_{1-3}alkyl$ or $C_{1-3}alkoxyC_{2-3}alkyl$) and R^{29} is as defined herein);
- 17) $C_{1-4}alkylX^9C_{1-4}alkylR^{28}$ (wherein X^9 and R^{28} are as defined herein);
- 18) $C_{2-3}alkenyl$ which may be unsubstituted or which may be substituted with one or more groups selected from hydroxy, fluoro, amino, $C_{1-4}alkylamino$, N,N -di($C_{1-4}alkyl$)amino, aminosulphonyl, N - $C_{1-4}alkylaminosulphonyl$ and N,N -di($C_{1-4}alkyl$)aminosulphonyl;

- 19) $C_{2,5}$ alkynyl which may be unsubstituted or which may be substituted with one or more groups selected from hydroxy, fluoro, amino, $C_{1,4}$ alkylamino, N,N -di($C_{1,4}$ alkyl)amino, aminosulphonyl, N - $C_{1,4}$ alkylaminosulphonyl and N,N -di($C_{1,4}$ alkyl)aminosulphonyl;
- 20) $C_{2,5}$ alkenyl $X^9C_{1,4}$ alkyl R^{28} (wherein X^9 and R^{28} are as defined herein);
- 5 21) $C_{2,5}$ alkynyl $X^9C_{1,4}$ alkyl R^{28} (wherein X^9 and R^{28} are as defined herein); and
- 22) $C_{1,4}$ alkyl $R^{54}(C_{1,4}alkyl)_q(X^9)_rR^{55}$ (wherein X^9 is as defined herein, q is 0 or 1, r is 0 or 1, and R^{54} and R^{55} are each independently selected from hydrogen, $C_{1,3}$ alkyl, cyclopentyl, cyclohexyl and a 5-6-membered saturated heterocyclic group with 1-2 heteroatoms, selected independently from O, S and N, which $C_{1,3}$ alkyl group may bear 1 or 2 substituents selected
- 10 from oxo, hydroxy, halogeno and $C_{1,4}$ alkoxy and which cyclic group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno, cyano, $C_{1,4}$ cyanoalkyl, $C_{1,4}$ alkyl, $C_{1,4}$ hydroxyalkyl, $C_{1,4}$ alkoxy, $C_{1,4}$ alkoxy $C_{1,4}$ alkyl, $C_{1,4}$ alkylsulphonyl $C_{1,4}$ alkyl, $C_{1,4}$ alkoxycarbonyl, $C_{1,4}$ aminoalkyl, $C_{1,4}$ alkylamino, di($C_{1,4}$ alkyl)amino, $C_{1,4}$ alkylamino $C_{1,4}$ alkyl, di($C_{1,4}$ alkyl)amino $C_{1,4}$ alkyl, $C_{1,4}$ alkylamino $C_{1,4}$ alkoxy, di($C_{1,4}$ alkyl)amino $C_{1,4}$ alkoxy and a
- 15 group $-(O-)_f(C_{1,4}alkyl)_g$ ringD (wherein f is 0 or 1, g is 0 or 1 and ring D is a 5-6-membered saturated heterocyclic group with 1-2 heteroatoms, selected independently from O, S and N, which cyclic group may bear one or more substituents selected from $C_{1,4}$ alkyl), with the proviso that R^{54} cannot be hydrogen);
- and additionally wherein any $C_{1,5}$ alkyl, $C_{2,5}$ alkenyl or $C_{2,5}$ alkynyl group in R^5X^1 may bear one
- 20 or more substituents selected from hydroxy, halogeno and amino);
- R^1 represents hydrogen, oxo, halogeno, hydroxy, $C_{1,4}$ alkoxy, $C_{1,4}$ alkyl, $C_{1,4}$ alkoxymethyl, $C_{1,4}$ alkanoyl, $C_{1,4}$ haloalkyl, cyano, amino, $C_{2,5}$ alkenyl, $C_{2,5}$ alkynyl, $C_{1,3}$ alkanoyloxy, nitro, $C_{1,4}$ alkanoylamino, $C_{1,4}$ alkoxycarbonyl, $C_{1,4}$ alkylsulphonyl, $C_{1,4}$ alkylsulphinyl, $C_{1,4}$ alkylsulphonyl, carbamoyl, N - $C_{1,4}$ alkylcarbamoyl, N,N -di($C_{1,4}$ alkyl)carbamoyl,
- 25 aminosulphonyl, N - $C_{1,4}$ alkylaminosulphonyl, N,N -di($C_{1,4}$ alkyl)aminosulphonyl, N -($C_{1,4}$ alkylsulphonyl)amino, N -($C_{1,4}$ alkylsulphonyl)- N -($C_{1,4}$ alkyl)amino, N,N -di($C_{1,4}$ alkylsulphonyl)amino, a $C_{3,7}$ alkylene chain joined to two ring C carbon atoms, $C_{1,4}$ alkanoylamino $C_{1,4}$ alkyl, carboxy or a group $R^{56}X^{10}$ (wherein X^{10} represents a direct bond, -O-, -CH₂-, -OC(O)-, -C(O)-, -S-, -SO-, -SO₂-, -NR⁵⁷C(O)-, -C(O)NR⁵⁸-, -SO₂NR⁵⁹-, -NR⁶⁰SO₂- or -
- 30 NR⁶¹- (wherein R^{57} , R^{58} , R^{59} , R^{60} and R^{61} each independently represents hydrogen, $C_{1,3}$ alkyl or $C_{1,3}$ alkoxy $C_{2,3}$ alkyl), and R^{56} is selected from one of the following twenty-two groups:

- 1) hydrogen, oxiranylC₁₋₄alkyl or C₁₋₃alkyl which may be unsubstituted or which may be substituted with one or more groups selected from hydroxy, fluoro, chloro, bromo and amino;
- 2) C₁₋₃alkylX¹¹C(O)R⁶² (wherein X¹¹ represents -O- or -NR⁶³- (in which R⁶³ represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl) and R⁶² represents C₁₋₃alkyl, -NR⁶⁴R⁶⁵ or -OR⁶⁶ (wherein R⁶⁴, R⁶⁵ and R⁶⁶ which may be the same or different each represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl));
- 3) C₁₋₃alkylX¹²R⁶⁷ (wherein X¹² represents -O-, -S-, -SO-, -SO₂-, -OC(O)-, -NR⁶⁸C(O)-, -C(O)NR⁶⁹-, -SO₂NR⁷⁰-, -NR⁷¹SO₂- or -NR⁷²- (wherein R⁶⁸, R⁶⁹, R⁷⁰, R⁷¹ and R⁷² each independently represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl) and R⁶⁷ represents hydrogen, C₁₋₃alkyl, cyclopentyl, cyclohexyl or a 5-6-membered saturated heterocyclic group with 1-2 heteroatoms, selected independently from O, S and N, which C₁₋₃alkyl group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno and C₁₋₄alkoxy and which cyclic group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno, cyano, C₁₋₄cyanoalkyl, C₁₋₄alkyl, C₁₋₄hydroxyalkyl, C₁₋₄alkoxy, C₁₋₄alkoxyC₁₋₄alkyl, C₁₋₄alkylsulphonylC₁₋₄alkyl, C₁₋₄alkoxycarbonyl, C₁₋₄aminoalkyl, C₁₋₄alkylamino, di(C₁₋₄alkyl)amino, C₁₋₄alkylaminoC₁₋₄alkyl, di(C₁₋₄alkyl)aminoC₁₋₄alkyl, C₁₋₄alkylaminoC₁₋₄alkoxy, di(C₁₋₄alkyl)aminoC₁₋₄alkoxy and a group $-(O-)_f(C_{1-4}alkyl)_gringD$ (wherein f is 0 or 1, g is 0 or 1 and ring D is a 5-6-membered saturated heterocyclic group with 1-2 heteroatoms, selected independently from O, S and N, which cyclic group may bear one or more substituents selected from C₁₋₄alkyl));
- 4) C₁₋₃alkylX¹³C₁₋₃alkylX¹⁴R⁷³ (wherein X¹³ and X¹⁴ which may be the same or different are each -O-, -S-, -SO-, -SO₂-, -NR⁷⁴C(O)-, -C(O)NR⁷⁵-, -SO₂NR⁷⁶-, -NR⁷⁷SO₂- or -NR⁷⁸- (wherein R⁷⁴, R⁷⁵, R⁷⁶, R⁷⁷ and R⁷⁸ each independently represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl) and R⁷³ represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl);
- 5) R⁷⁹ (wherein R⁷⁹ is a 5-6-membered saturated heterocyclic group (linked via carbon or nitrogen) with 1-2 heteroatoms, selected independently from O, S and N, which heterocyclic group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno, cyano, C₁₋₄cyanoalkyl, C₁₋₄alkyl, C₁₋₄hydroxyalkyl, C₁₋₄alkoxy, C₁₋₄alkoxyC₁₋₄alkyl, C₁₋₄alkylsulphonylC₁₋₄alkyl, C₁₋₄alkoxycarbonyl, C₁₋₄aminoalkyl, C₁₋₄alkylamino, di(C₁₋₄alkyl)amino, C₁₋₄alkylaminoC₁₋₄alkyl, di(C₁₋₄alkyl)aminoC₁₋₄alkyl, C₁₋₄alkylaminoC₁₋₄alkoxy, di(C₁₋₄alkyl)aminoC₁₋₄alkoxy and a group $-(O-)_f(C_{1-4}alkyl)_gringD$ (wherein f is 0 or 1, g is 0 or 1 and ring D is a 5-6-membered saturated heterocyclic group with 1-2 heteroatoms, selected

independently from O, S and N, which cyclic group may bear one or more substituents selected from C₁₋₄alkyl));

6) C₁₋₃alkylR⁷⁹ (wherein R⁷⁹ is as defined herein);

7) C₂₋₃alkenylR⁷⁹ (wherein R⁷⁹ is as defined herein);

5 8) C₂₋₃alkynylR⁷⁹ (wherein R⁷⁹ is as defined herein);

9) R⁸⁰ (wherein R⁸⁰ represents a pyridone group, a phenyl group or a 5-6-membered aromatic heterocyclic group (linked via carbon or nitrogen) with 1-3 heteroatoms selected from O, N and S, which pyridone, phenyl or aromatic heterocyclic group may carry up to 5 substituents selected from hydroxy, halogeno, amino, C₁₋₄alkyl, C₁₋₄alkoxy, C₁₋₄hydroxyalkyl, C₁₋

10 4aminoalkyl, C₁₋₄alkylamino, C₁₋₄hydroxyalkoxy, carboxy, trifluoromethyl, cyano, -C(O)NR⁸¹R⁸², -NR⁸³C(O)R⁸⁴ (wherein R⁸¹, R⁸², R⁸³ and R⁸⁴, which may be the same or different, each represents hydrogen, C₁₋₄alkyl or C₁₋₃alkoxyC₂₋₃alkyl) and a group -(O-)_f(C₁₋₄alkyl)_gringD (wherein f is 0 or 1, g is 0 or 1 and ring D is a 5-6-membered saturated heterocyclic group with 1-2 heteroatoms, selected independently from O, S and N, which
15 cyclic group may bear one or more substituents selected from C₁₋₄alkyl));

10) C₁₋₃alkylR⁸⁰ (wherein R⁸⁰ is as defined herein);

11) C₂₋₃alkenylR⁸⁰ (wherein R⁸⁰ is as defined herein);

12) C₂₋₃alkynylR⁸⁰ (wherein R⁸⁰ is as defined herein);

13) C₁₋₃alkylX¹⁵R⁸⁰ (wherein X¹⁵ represents -O-, -S-, -SO-, -SO₂-, -NR⁸⁵C(O)-, -C(O)NR⁸⁶-, -
20 SO₂NR⁸⁷-, -NR⁸⁸SO₂- or -NR⁸⁹- (wherein R⁸⁵, R⁸⁶, R⁸⁷, R⁸⁸ and R⁸⁹ each independently represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl) and R⁸⁰ is as defined herein);

14) C₂₋₃alkenylX¹⁶R⁸⁰ (wherein X¹⁶ represents -O-, -S-, -SO-, -SO₂-, -NR⁹⁰C(O)-, -C(O)NR⁹¹-,
-SO₂NR⁹²-, -NR⁹³SO₂- or -NR⁹⁴- (wherein R⁹⁰, R⁹¹, R⁹², R⁹³ and R⁹⁴ each independently represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl) and R⁸⁰ is as defined herein);

25 15) C₂₋₃alkynylX¹⁷R⁸⁰ (wherein X¹⁷ represents -O-, -S-, -SO-, -SO₂-, -NR⁹⁵C(O)-, -C(O)NR⁹⁶-,
-SO₂NR⁹⁷-, -NR⁹⁸SO₂- or -NR⁹⁹- (wherein R⁹⁵, R⁹⁶, R⁹⁷, R⁹⁸ and R⁹⁹ each independently represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl) and R⁸⁰ is as defined herein);

16) C₁₋₄alkylX¹⁸C₁₋₄alkylR⁸⁰ (wherein X¹⁸ represents -O-, -S-, -SO-, -SO₂-, -NR¹⁰⁰C(O)-, -
30 C(O)NR¹⁰¹-, -SO₂NR¹⁰²-, -NR¹⁰³SO₂- or -NR¹⁰⁴- (wherein R¹⁰⁰, R¹⁰¹, R¹⁰², R¹⁰³ and R¹⁰⁴ each independently represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl) and R⁸⁰ is as defined herein);

17) C₁₋₄alkylX¹⁸C₁₋₄alkylR⁷⁹ (wherein X¹⁸ and R⁷⁹ are as defined herein);

- 321 -

18) $C_{2,5}$ alkenyl which may be unsubstituted or which may be substituted with one or more groups selected from hydroxy, fluoro, amino, $C_{1,4}$ alkylamino, N,N -di($C_{1,4}$ alkyl)amino, aminosulphonyl, N - $C_{1,4}$ alkylaminosulphonyl and N,N -di($C_{1,4}$ alkyl)aminosulphonyl;

19) $C_{2,5}$ alkynyl which may be unsubstituted or which may be substituted with one or more groups selected from hydroxy, fluoro, amino, $C_{1,4}$ alkylamino, N,N -di($C_{1,4}$ alkyl)amino, aminosulphonyl, N - $C_{1,4}$ alkylaminosulphonyl and N,N -di($C_{1,4}$ alkyl)aminosulphonyl;

20) $C_{2,5}$ alkenyl $X^{18}C_{1,4}$ alkyl R^{79} (wherein X^{18} and R^{79} are as defined herein);

21) $C_{2,5}$ alkynyl $X^{18}C_{1,4}$ alkyl R^{79} (wherein X^{18} and R^{79} are as defined herein); and

22) $C_{1,4}$ alkyl $R^{105}(C_{1,4}alkyl)_x(X^{18})_yR^{106}$ (wherein X^{18} is as defined herein, x is 0 or 1, y is 0 or 1,

and R^{105} and R^{106} are each independently selected from hydrogen, $C_{1,3}$ alkyl, cyclopentyl, cyclohexyl and a 5-6-membered saturated heterocyclic group with 1-2 heteroatoms, selected independently from O, S and N, which $C_{1,3}$ alkyl group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno and $C_{1,4}$ alkoxy and which cyclic group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno, cyano, $C_{1,4}$ cyanoalkyl, $C_{1,4}$ alkyl, $C_{1,4}$ hydroxyalkyl, $C_{1,4}$ alkoxy, $C_{1,4}$ alkoxy $C_{1,4}$ alkyl, $C_{1,4}$ alkylsulphonyl $C_{1,4}$ alkyl, $C_{1,4}$ alkoxycarbonyl, $C_{1,4}$ aminoalkyl, $C_{1,4}$ alkylamino, di($C_{1,4}$ alkyl)amino, $C_{1,4}$ alkylamino $C_{1,4}$ alkyl, di($C_{1,4}$ alkyl)amino $C_{1,4}$ alkyl, $C_{1,4}$ alkylamino $C_{1,4}$ alkoxy, di($C_{1,4}$ alkyl)amino $C_{1,4}$ alkoxy and a group $-(O-)_f(C_{1,4}alkyl)_g$ ringD (wherein f is 0 or 1, g is 0 or 1 and ring D is a 5-6-membered saturated heterocyclic group with 1-2 heteroatoms, selected independently from O, S and N,

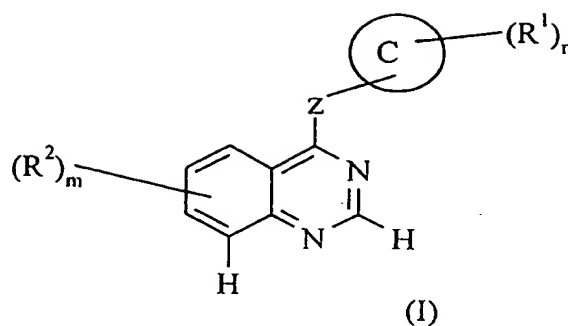
which cyclic group may bear one or more substituents selected from $C_{1,4}$ alkyl) with the proviso that R^{105} cannot be hydrogen);

and additionally wherein any $C_{1,3}$ alkyl, $C_{2,5}$ alkenyl or $C_{2,5}$ alkynyl group in $R^{56}X^{10}$ may bear one or more substituents selected from hydroxy, halogeno and amino);

or a salt thereof in the manufacture of a medicament for use in the production of an

antiangiogenic and/or vascular permeability reducing effect in warm-blooded animals such as humans.

2. The use of a compound of the formula according to claim 1:



wherein:

ring C is a 9-10-membered bicyclic moiety which may be saturated or unsaturated, which may be aromatic or non-aromatic, and which optionally may contain 1-3 heteroatoms selected independently from O, N and S;

Z is -O-, -NH-, -S-, -CH₂- or a direct bond;

R¹ represents hydrogen, oxo, halogeno, hydroxy, C₁₋₄alkoxy, C₁₋₄alkyl, C₁₋₄alkoxymethyl, C₁₋₄alkanoyl, C₁₋₄haloalkyl, cyano, amino, C₂₋₅alkenyl, C₂₋₅alkynyl, C₁₋₃alkanoyloxy, nitro, C₁₋₄alkanoylamino, C₁₋₄alkoxycarbonyl, C₁₋₄alkylsulphanyl, C₁₋₄alkylsulphinyl, C₁₋₄alkylsulphonyl, carbamoyl, N-C₁₋₄alkylcarbamoyl, N,N-di(C₁₋₄alkyl)carbamoyl, aminosulphonyl, N-C₁₋₄alkylaminosulphonyl, N,N-di(C₁₋₄alkyl)aminosulphonyl, N-(C₁₋₄alkylsulphonyl)amino, N-(C₁₋₄alkylsulphonyl)-N-(C₁₋₄alkyl)amino, N,N-di(C₁₋₄alkylsulphonyl)amino or a C₃₋₇alkylene chain joined to two ring C carbon atoms;

n is an integer from 0 to 5;

m is an integer from 0 to 3;

R² represents hydrogen, hydroxy, halogeno, cyano, nitro, trifluoromethyl, C₁₋₃alkyl, C₁₋₃alkoxy, C₁₋₃alkylsulphanyl, -NR³R⁴ (wherein R³ and R⁴, which may be the same or different, each represents hydrogen or C₁₋₃alkyl), or R⁵X¹- (wherein X¹ represents a direct bond, -O-, -CH₂-, -OC(O)-, -C(O)-, -S-, -SO-, -SO₂-, -NR⁶C(O)-, -C(O)NR⁷-, -SO₂NR⁸-, -NR⁹SO₂- or -NR¹⁰- (wherein R⁶, R⁷, R⁸, R⁹ and R¹⁰ each independently represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl), and R⁵ is selected from one of the following twenty-one groups:

1) hydrogen or C₁₋₃alkyl which may be unsubstituted or which may be substituted with one or more groups selected from hydroxy, fluoro and amino;

2) C₁₋₃alkylX²C(O)R¹¹ (wherein X² represents -O- or -NR¹²- (in which R¹² represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl) and R¹¹ represents C₁₋₃alkyl, -NR¹³R¹⁴ or -OR¹⁵

- 323 -

(wherein R^{13} , R^{14} and R^{15} which may be the same or different each represents hydrogen, C_{1-3} alkyl or C_{1-3} alkoxy C_{2-3} alkyl));

3) C_{1-3} alkyl X^3R^{16} (wherein X^3 represents -O-, -S-, -SO-, -SO₂-, -OC(O)-, -NR¹⁷C(O)-, -C(O)NR¹⁸-, -SO₂NR¹⁹-, -NR²⁰SO₂- or -NR²¹- (wherein R^{17} , R^{18} , R^{19} , R^{20} and R^{21} each

5 independently represents hydrogen, C_{1-3} alkyl or C_{1-3} alkoxy C_{2-3} alkyl) and R^{16} represents hydrogen, C_{1-3} alkyl, cyclopentyl, cyclohexyl or a 5-6-membered saturated heterocyclic group with 1-2 heteroatoms, selected independently from O, S and N, which C_{1-3} alkyl group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno and C_{1-4} alkoxy and which cyclic group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno, C_{1-4} alkyl, C_{1-4} hydroxyalkyl and C_{1-4} alkoxy);

4) C_{1-3} alkyl X^4C_{1-3} alkyl X^5R^{22} (wherein X^4 and X^5 which may be the same or different are each -O-, -S-, -SO-, -SO₂-, -NR²³C(O)-, -C(O)NR²⁴-, -SO₂NR²⁵-, -NR²⁶SO₂- or -NR²⁷- (wherein R^{23} , R^{24} , R^{25} , R^{26} and R^{27} each independently represents hydrogen, C_{1-3} alkyl or C_{1-3} alkoxy C_{2-3} alkyl) and R^{22} represents hydrogen or C_{1-3} alkyl);

5) R^{28} (wherein R^{28} is a 5-6-membered saturated heterocyclic group (linked via carbon or nitrogen) with 1-2 heteroatoms, selected independently from O, S and N, which heterocyclic group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno, cyano, C_{1-4} cyanoalkyl, C_{1-4} alkyl, C_{1-4} hydroxyalkyl, C_{1-4} alkoxy, C_{1-4} alkoxy C_{1-4} alkyl and C_{1-4} alkylsulphonyl C_{1-4} alkyl);

6) C_{1-5} alkyl R^{28} (wherein R^{28} is as defined herein);

7) C_{2-5} alkenyl R^{28} (wherein R^{28} is as defined herein);

8) C_{2-5} alkynyl R^{28} (wherein R^{28} is as defined herein);

9) R^{29} (wherein R^{29} represents a pyridone group, a phenyl group or a 5-6-membered aromatic heterocyclic group (linked via carbon or nitrogen) with 1-3 heteroatoms selected from O, N and S, which pyridone, phenyl or aromatic heterocyclic group may carry up to 5 substituents on an available carbon atom selected from hydroxy, halogeno, amino, C_{1-4} alkyl, C_{1-4} alkoxy, C_{1-4} hydroxyalkyl, C_{1-4} aminoalkyl, C_{1-4} alkylamino, C_{1-4} hydroxyalkoxy, carboxy, trifluoromethyl, cyano, -C(O)NR³⁰R³¹ and -NR³²C(O)R³³ (wherein R^{30} , R^{31} , R^{32} and R^{33} , which may be the same or different, each represents hydrogen, C_{1-4} alkyl or C_{1-3} alkoxy C_{2-3} alkyl));

10) C_{1-5} alkyl R^{29} (wherein R^{29} is as defined herein);

11) C_{2-5} alkenyl R^{29} (wherein R^{29} is as defined herein);

12) C_{2-5} alkynyl R^{29} (wherein R^{29} is as defined herein);

- 324 -

- 13) $C_{1-3}alkylX^6R^{29}$ (wherein X^6 represents -O-, -S-, -SO-, -SO₂-, -NR³⁴C(O)-, -C(O)NR³⁵-, -SO₂NR³⁶-, -NR³⁷SO₂- or -NR³⁸- (wherein R³⁴, R³⁵, R³⁶, R³⁷ and R³⁸ each independently represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl) and R²⁹ is as defined herein);
- 14) $C_{2-5}alkenylX^7R^{29}$ (wherein X^7 represents -O-, -S-, -SO-, -SO₂-, -NR³⁹C(O)-, -C(O)NR⁴⁰-, -SO₂NR⁴¹-, -NR⁴²SO₂- or -NR⁴³- (wherein R³⁹, R⁴⁰, R⁴¹, R⁴² and R⁴³ each independently represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl) and R²⁹ is as defined herein);
- 15) $C_{2-5}alkynylX^8R^{29}$ (wherein X^8 represents -O-, -S-, -SO-, -SO₂-, -NR⁴⁴C(O)-, -C(O)NR⁴⁵-, -SO₂NR⁴⁶-, -NR⁴⁷SO₂- or -NR⁴⁸- (wherein R⁴⁴, R⁴⁵, R⁴⁶, R⁴⁷ and R⁴⁸ each independently represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl) and R²⁹ is as defined herein);
- 16) $C_{1-3}alkylX^9C_{1-3}alkylR^{29}$ (wherein X^9 represents -O-, -S-, -SO-, -SO₂-, -NR⁴⁹C(O)-, -C(O)NR⁵⁰-, -SO₂NR⁵¹-, -NR⁵²SO₂- or -NR⁵³- (wherein R⁴⁹, R⁵⁰, R⁵¹, R⁵² and R⁵³ each independently represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl) and R²⁹ is as defined herein);
- 17) $C_{1-3}alkylX^9C_{1-3}alkylR^{28}$ (wherein X^9 and R²⁸ are as defined herein);
- 18) $C_{2-5}alkenyl$ which may be unsubstituted or which may be substituted with one or more groups selected from hydroxy, fluoro, amino, C₁₋₄alkylamino, N,N-di(C₁₋₄alkyl)amino, aminosulphonyl, N-C₁₋₄alkylaminosulphonyl and N,N-di(C₁₋₄alkyl)aminosulphonyl;
- 19) $C_{2-5}alkynyl$ which may be unsubstituted or which may be substituted with one or more groups selected from hydroxy, fluoro, amino, C₁₋₄alkylamino, N,N-di(C₁₋₄alkyl)amino, aminosulphonyl, N-C₁₋₄alkylaminosulphonyl and N,N-di(C₁₋₄alkyl)aminosulphonyl;
- 20) $C_{2-5}alkenylX^9C_{1-4}alkylR^{28}$ (wherein X^9 and R²⁸ are as defined herein); and
- 21) $C_{2-5}alkynylX^9C_{1-4}alkylR^{28}$ (wherein X^9 and R²⁸ are as defined herein);
- and salts thereof, and prodrugs thereof for example esters, amides and sulphides, in the manufacture of a medicament for use in the production of an antiangiogenic and/or vascular permeability reducing effect in warm-blooded animals such as humans.

3. The use of a compound of the formula I according to claim 1, wherein R² represents hydroxy, halogeno, cyano, nitro, trifluoromethyl, C₁₋₃alkyl, amino or R⁵X¹- [wherein X¹ is as defined in claim 1 and R⁵ is selected from one of the following twenty-two groups:

- 1) C₁₋₄alkyl which may be unsubstituted or which may be substituted with one or more groups selected from fluoro, chloro and bromo, or C₂₋₃alkyl which may be unsubstituted or substituted with one or more groups selected from hydroxy and amino;
- 2) C₂₋₃alkylX²C(O)R¹¹ (wherein X² is as defined in claim 1 and R¹¹ represents -NR¹³R¹⁴ or -OR¹⁵ (wherein R¹³, R¹⁴ and R¹⁵ which may be the same or different are each C₁₋₄alkyl or C₁₋₃alkoxyethyl));
- 3) C₂₋₄alkylX³R¹⁶ (wherein X³ is as defined in claim 1 and R¹⁶ is a group selected from C₁₋₃alkyl, cyclopentyl, cyclohexyl, pyrrolidinyl, piperazinyl, piperidinyl, imidazolidinyl, azetidiny and tetrahydropyranyl, which C₁₋₃alkyl group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno and C₁₋₂alkoxy and which cyclopentyl, cyclohexyl, pyrrolidinyl, piperazinyl, piperidinyl, imidazolidinyl, azetidiny or tetrahydropyranyl group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno, cyano, C₁₋₃cyanoalkyl, C₁₋₃alkyl, C₁₋₃hydroxyalkyl, C₁₋₃alkoxy, C₁₋₂alkoxyC₁₋₃alkyl, C₁₋₂alkylsulphonylC₁₋₃alkyl, C₁₋₃alkoxycarbonyl, C₁₋₃alkylamino, di(C₁₋₃alkyl)amino, C₁₋₃alkylaminoC₁₋₃alkyl, di(C₁₋₃alkyl)aminoC₁₋₃alkyl, C₁₋₃alkylaminoC₁₋₃alkoxy, di(C₁₋₃alkyl)aminoC₁₋₃alkoxy and a group -(-O-)_f(C₁₋₃alkyl)_gringD (wherein f is 0 or 1, g is 0 or 1 and ring D is a heterocyclic group selected from pyrrolidinyl, piperazinyl, piperidinyl, imidazolidinyl, azetidiny, morpholino and thiomorpholino, which cyclic group may bear one or more substituents selected from C₁₋₃alkyl));
- 4) C₂₋₃alkylX⁴C₂₋₃alkylX⁵R²² (wherein X⁴ and X⁵ are as defined in claim 1 and R²² represents hydrogen or C₁₋₃alkyl);
- 5) R²⁸ (wherein R²⁸ is as defined in claim 1);
- 6) C₁₋₄alkylR¹¹⁰ (wherein R¹¹⁰ is a group selected from pyrrolidinyl, piperazinyl, piperidinyl, imidazolidin-1-yl, azetidiny, 1,3-dioxolan-2-yl, 1,3-dioxan-2-yl, 1,3-dithiolan-2-yl and 1,3-dithian-2-yl, which group is linked to C₁₋₄alkyl through a carbon atom and which group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno, cyano, C₁₋₃cyanoalkyl, C₁₋₃alkyl, C₁₋₃hydroxyalkyl, C₁₋₃alkoxy, C₁₋₂alkoxyC₁₋₃alkyl, C₁₋₂alkylsulphonylC₁₋₃alkyl, C₁₋₃alkoxycarbonyl, C₁₋₃alkylamino, di(C₁₋₃alkyl)amino, C₁₋₃alkylaminoC₁₋₃alkyl, di(C₁₋₃alkyl)aminoC₁₋₃alkyl, C₁₋₃alkylaminoC₁₋₃alkoxy, di(C₁₋₃alkyl)aminoC₁₋₃alkoxy and a group -(-O-)_f(C₁₋₃alkyl)_gringD (wherein f is 0 or 1, g is 0 or 1 and ring D is a heterocyclic group selected from pyrrolidinyl, piperazinyl, piperidinyl, imidazolidinyl, azetidiny, morpholino and thiomorpholino, which cyclic group may bear one or more substituents selected from C₁₋

- alkyl)) or C₂₋₄alkylR¹¹¹ (wherein R¹¹¹ is a group selected from morpholino, thiomorpholino, azetidin-1-yl, pyrrolidin-1-yl, piperazin-1-yl and piperidino which group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno, cyano, C₁₋₃cyanoalkyl, C₁₋₃alkyl, C₁₋₃hydroxyalkyl, C₁₋₃alkoxy, C₁₋₂alkoxyC₁₋₃alkyl, C₁₋₂alkylsulphonylC₁₋₃alkyl, C₁₋₃alkoxycarbonyl, C₁₋₃alkylamino, di(C₁₋₃alkyl)amino, C₁₋₃alkylaminoC₁₋₃alkyl, di(C₁₋₃alkyl)aminoC₁₋₃alkyl, C₁₋₃alkylaminoC₁₋₃alkoxy, di(C₁₋₃alkyl)aminoC₁₋₃alkoxy and a group -(-O-)_f(C₁₋₃alkyl)_gringD (wherein f is 0 or 1, g is 0 or 1 and ring D is a heterocyclic group selected from pyrrolidinyl, piperazinyl, piperidinyl, imidazolidinyl, azetidinyl, morpholino and thiomorpholino, which cyclic group may bear one or more substituents selected from C₁₋₃alkyl));
- 7) C₃₋₄alkenylR¹¹² (wherein R¹¹² represents R¹¹⁰ or R¹¹¹ as defined herein);
- 8) C₃₋₄alkynylR¹¹² (wherein R¹¹² represents R¹¹⁰ or R¹¹¹ as defined herein);
- 9) R²⁹ (wherein R²⁹ is as defined in claim 1);
- 10) C₁₋₄alkylR²⁹ (wherein R²⁹ is as defined in claim 1);
- 11) 1-R²⁹prop-1-en-3-yl or 1-R²⁹but-2-en-4-yl (wherein R²⁹ is as defined in claim 1 with the proviso that when R⁵ is 1-R²⁹prop-1-en-3-yl, R²⁹ is linked to the alkenyl group via a carbon atom);
- 12) 1-R²⁹prop-1-yn-3-yl or 1-R²⁹but-2-yn-4-yl (wherein R²⁹ is as defined in claim 1 with the proviso that when R⁵ is 1-R²⁹prop-1-yn-3-yl, R²⁹ is linked to the alkynyl group via a carbon atom);
- 13) C₁₋₅alkylX⁶R²⁹ (wherein X⁶ and R²⁹ are as defined in claim 1);
- 14) 1-(R²⁹X⁷)but-2-en-4-yl (wherein X⁷ and R²⁹ are as defined in claim 1);
- 15) 1-(R²⁹X⁸)but-2-yn-4-yl (wherein X⁸ and R²⁹ are as defined in claim 1);
- 16) C₂₋₃alkylX⁹C₁₋₃alkylR²⁹ (wherein X⁹ and R²⁹ are as defined in claim 1);
- 17) C₂₋₃alkylX⁹C₁₋₃alkylR²⁸ (wherein X⁹ and R²⁸ are as defined in claim 1);
- 18) C₂₋₅alkenyl which may be unsubstituted or which may be substituted with one or more fluorine atoms or with one or two groups selected from hydroxy, fluoro, amino, C₁₋₄alkylamino, N,N-di(C₁₋₄alkyl)amino, aminosulphonyl, N-C₁₋₄alkylaminosulphonyl and N,N-di(C₁₋₄alkyl)aminosulphonyl;
- 19) C₂₋₅alkynyl which may be unsubstituted or which may be substituted with one or more fluorine atoms or with one or two groups selected from hydroxy, fluoro, amino, C₁₋

alkylamino, N,N-di(C₁₋₄alkyl)amino, aminosulphonyl, N-C₁₋₄alkylaminosulphonyl and N,N-di(C₁₋₄alkyl)aminosulphonyl;

20) C₂₋₄alkenylX⁹C₁₋₃alkylR²⁸ (wherein X⁹ and R²⁸ are as defined in claim 1);

21) C₂₋₄alkynylX⁹C₁₋₃alkylR²⁸ (wherein X⁹ and R²⁸ are as defined in claim 1); and

5 22) C₁₋₃alkylR⁵⁴(C₁₋₃alkyl)_q(X⁹)_rR⁵⁵ (wherein X⁹, q, r, R⁵⁴ and R⁵⁵ are as defined in claim 1);

and additionally wherein any C₁₋₃alkyl, C₂₋₅alkenyl or C₂₋₅alkynyl group in R⁵X¹- may bear one or more substituents selected from hydroxy, halogeno and amino].

10 4. The use of a compound of the formula I according to any one of the preceding claims wherein Z is -O-, -NH- or -S-.

15 5. The use of a compound of the formula I according to any one of the preceding claims wherein ring C is a 9-10-membered heteroaromatic bicyclic moiety which contains 1-3 heteroatoms selected independently from O, N and S.

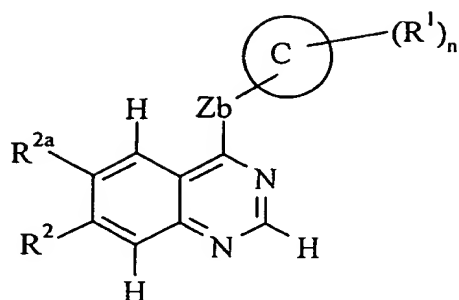
20 6. The use of a compound of the formula I according to any one of the preceding claims wherein R¹ represents oxo, halogeno, hydroxy, C₁₋₂alkoxy, C₁₋₂alkyl, C₁₋₂alkoxymethyl, C₂₋₃alkanoyl, C₁₋₂haloalkyl, cyano, amino, C₂₋₄alkenyl, C₂₋₄alkynyl, C₂₋₃alkanoyloxy, nitro, C₂₋₃alkanoylamino, C₁₋₂alkoxycarbonyl, C₁₋₂alkylsulphanyl, C₁₋₂alkylsulphinyl, C₁₋₂alkylsulphonyl, carbamoyl, N-C₁₋₂alkylcarbamoyl, N,N-di(C₁₋₂alkyl)carbamoyl, aminosulphonyl, N-C₁₋₂alkylaminosulphonyl, N,N-di(C₁₋₂alkyl)aminosulphonyl, N-(C₁₋₂alkylsulphonyl)amino, N-(C₁₋₂alkylsulphonyl)-N-(C₁₋₂alkyl)amino or a C₃₋₇alkylene chain joined to two ring C carbon atoms.

25 7. The use of a compound of the formula I according to any one of the preceding claims wherein n is 0, 1 or 2.

30 8. The use of a compound of the formula I according to any one of the preceding claims wherein m is 1 or 2.

9. A compound of the formula II:

- 328 -



(II)

- 10 [wherein:
 ring C, R^1 , R^2 and n are as defined in claim 1, Zb is -O- or -S- and R^{2a} represents hydrogen,
 halogeno, C_{1-3} alkyl, trifluoromethyl, C_{1-3} alkoxy, C_{1-3} alkylsulphanyl, $-NR^{3a}R^{4a}$ (wherein R^{3a} and
 R^{4a} , which may be the same or different, each represents hydrogen or C_{1-3} alkyl), or
 $R^{5a}(CH_2)_{za}X^{1a}$ (wherein R^{5a} is a 5- or 6-membered saturated heterocyclic group with 1-2
 15 heteroatoms, selected independently from O, S and N, which heterocyclic group may bear 1 or
 2 substituents selected from oxo, hydroxy, halogeno, cyano, C_{1-4} cyanoalkyl, C_{1-4} alkyl, C_{1-4}
 hydroxyalkyl, C_{1-4} alkoxy, C_{1-4} alkoxy C_{1-4} alkyl, C_{1-4} alkylsulphonyl C_{1-4} alkyl, C_{1-4}
 alkoxy carbonyl, C_{1-4} aminoalkyl, C_{1-4} alkylamino, di(C_{1-4} alkyl)amino, C_{1-4} alkylamino C_{1-4} alkyl,
 di(C_{1-4} alkyl)amino C_{1-4} alkyl, C_{1-4} alkylamino C_{1-4} alkoxy, di(C_{1-4} alkyl)amino C_{1-4} alkoxy and a
 20 group $-(O-)_f(C_{1-4}alkyl)_g$ ringD (wherein f is 0 or 1, g is 0 or 1 and ring D is a 5-6-membered
 saturated heterocyclic group with 1-2 heteroatoms, selected independently from O, S and N,
 which cyclic group may bear one or more substituents selected from C_{1-4} alkyl), za is an
 integer from 0 to 4 and X^{1a} represents a direct bond, -O-, $-CH_2-$, -S-, -SO-, $-SO_2-$, $-NR^{6a}C(O)-$,
 $-C(O)NR^{7a}-$, $-SO_2NR^{8a}-$, $-NR^{9a}SO_2-$ or $-NR^{10a}-$ (wherein R^{6a} , R^{7a} , R^{8a} , R^{9a} and R^{10a} each
 25 independently represents hydrogen, C_{1-3} alkyl or C_{1-3} alkoxy C_{2-3} alkyl));
 with the proviso that R^2 is not hydrogen and excluding the compounds:
 6,7-dimethoxy-4-(1-naphthylsulphanyl)quinazoline, 6,7-dimethoxy-4-(2-
 naphthylsulphanyl)quinazoline, 6,7-dimethoxy-4-(1-naphthyloxy)quinazoline and 6,7-
 dimethoxy-4-(2-naphthyloxy)quinazoline;
 30 or a salt thereof.

10. A compound of the formula II according to claim 9 wherein R² represents hydroxy, halogeno, cyano, nitro, trifluoromethyl, C₁₋₃alkyl, amino or R⁵X¹- [wherein X¹ is as defined in claim 1 and R⁵ is selected from one of the following twenty-two groups:

- 1) C₁₋₄alkyl which may be unsubstituted or which may be substituted with one or more groups selected from fluoro, chloro and bromo, or C₂₋₅alkyl which may be unsubstituted or substituted with one or more groups selected from hydroxy and amino;
- 2) C₂₋₃alkylX²C(O)R¹¹ (wherein X² is as defined in claim 1 and R¹¹ represents -NR¹³R¹⁴ or -OR¹⁵ (wherein R¹³, R¹⁴ and R¹⁵ which may be the same or different are each C₁₋₄alkyl or C₁₋₂alkoxyethyl));
- 3) C₂₋₄alkylX³R¹⁶ (wherein X³ is as defined in claim 1 and R¹⁶ is a group selected from C₁₋₃alkyl, cyclopentyl, cyclohexyl, pyrrolidinyl, piperazinyl, piperidinyl, imidazolidinyl, azetidiny and tetrahydropyranyl, which C₁₋₃alkyl group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno and C₁₋₂alkoxy and which cyclopentyl, cyclohexyl, pyrrolidinyl, piperazinyl, piperidinyl, imidazolidinyl, azetidiny or tetrahydropyranyl group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno, cyano, C₁₋₃cyanoalkyl, C₁₋₃alkyl, C₁₋₃hydroxyalkyl, C₁₋₃alkoxy, C₁₋₂alkoxyC₁₋₃alkyl, C₁₋₂alkylsulphonylC₁₋₃alkyl, C₁₋₃alkoxycarbonyl, C₁₋₃alkylamino, di(C₁₋₃alkyl)amino, C₁₋₃alkylaminoC₁₋₃alkyl, di(C₁₋₃alkyl)aminoC₁₋₃alkyl, C₁₋₃alkylaminoC₁₋₃alkoxy, di(C₁₋₃alkyl)aminoC₁₋₃alkoxy and a group -(-O-)_f(C₁₋₃alkyl)_gringD (wherein f is 0 or 1, g is 0 or 1 and ring D is a heterocyclic group selected from pyrrolidinyl, piperazinyl, piperidinyl, imidazolidinyl, azetidiny, morpholino and thiomorpholino, which cyclic group may bear one or more substituents selected from C₁₋₃alkyl));
- 4) C₂₋₃alkylX⁴C₂₋₃alkylX⁵R²² (wherein X⁴ and X⁵ are as defined in claim 1 and R²² represents hydrogen or C₁₋₃alkyl);
- 5) R²⁸ (wherein R²⁸ is as defined in claim 1);
- 6) C₁₋₄alkylR¹¹⁰ (wherein R¹¹⁰ is a group selected from pyrrolidinyl, piperazinyl, piperidinyl, imidazolidin-1-yl, azetidiny, 1,3-dioxolan-2-yl, 1,3-dioxan-2-yl, 1,3-dithiolan-2-yl and 1,3-dithian-2-yl, which group is linked to C₁₋₄alkyl through a carbon atom and which group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno, cyano, C₁₋₃cyanoalkyl, C₁₋₃alkyl, C₁₋₃hydroxyalkyl, C₁₋₃alkoxy, C₁₋₂alkoxyC₁₋₃alkyl, C₁₋₂alkylsulphonylC₁₋₃alkyl, C₁₋₃alkoxycarbonyl, C₁₋₃alkylamino, di(C₁₋₃alkyl)amino, C₁₋₃alkylaminoC₁₋₃alkyl, di(C₁₋₃alkyl)aminoC₁₋₃alkyl, C₁₋₃alkylaminoC₁₋₃alkoxy, di(C₁₋₃alkyl)aminoC₁₋₃alkoxy and a group -(-

O-)_f(C₁₋₃alkyl)_gringD (wherein f is 0 or 1, g is 0 or 1 and ring D is a heterocyclic group selected from pyrrolidinyl, piperazinyl, piperidinyl, imidazolidinyl, azetidiny, morpholino and thiomorpholino, which cyclic group may bear one or more substituents selected from C₁₋₃alkyl)) or C₂₋₄alkylR¹¹¹ (wherein R¹¹¹ is a group selected from morpholino, thiomorpholino, azetidin-1-yl, pyrrolidin-1-yl, piperazin-1-yl and piperidino which group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno, cyano, C₁₋₃cyanoalkyl, C₁₋₃alkyl, C₁₋₃hydroxyalkyl, C₁₋₃alkoxy, C₁₋₂alkoxyC₁₋₃alkyl, C₁₋₂alkylsulphonylC₁₋₃alkyl, C₁₋₃alkoxycarbonyl, C₁₋₃alkylamino, di(C₁₋₃alkyl)amino, C₁₋₃alkylaminoC₁₋₃alkyl, di(C₁₋₃alkyl)aminoC₁₋₃alkyl, C₁₋₃alkylaminoC₁₋₃alkoxy, di(C₁₋₃alkyl)aminoC₁₋₃alkoxy and a group -(-

O-)_f(C₁₋₃alkyl)_gringD (wherein f is 0 or 1, g is 0 or 1 and ring D is a heterocyclic group selected from pyrrolidinyl, piperazinyl, piperidinyl, imidazolidinyl, azetidiny, morpholino and thiomorpholino, which cyclic group may bear one or more substituents selected from C₁₋₃alkyl));

7) C₃₋₄alkenylR¹¹² (wherein R¹¹² represents R¹¹⁰ or R¹¹¹ as defined herein);

8) C₃₋₄alkynylR¹¹² (wherein R¹¹² represents R¹¹⁰ or R¹¹¹ as defined herein);

9) R²⁹ (wherein R²⁹ is as defined in claim 1);

10) C₁₋₄alkylR²⁹ (wherein R²⁹ is as defined in claim 1);

11) 1-R²⁹prop-1-en-3-yl or 1-R²⁹but-2-en-4-yl (wherein R²⁹ is as defined in claim 1 with the proviso that when R⁵ is 1-R²⁹prop-1-en-3-yl, R²⁹ is linked to the alkenyl group via a carbon atom);

12) 1-R²⁹prop-1-yn-3-yl or 1-R²⁹but-2-yn-4-yl (wherein R²⁹ is as defined in claim 1 with the proviso that when R⁵ is 1-R²⁹prop-1-yn-3-yl, R²⁹ is linked to the alkynyl group via a carbon atom);

13) C₁₋₅alkylX⁶R²⁹ (wherein X⁶ and R²⁹ are as defined in claim 1);

14) 1-(R²⁹X⁷)but-2-en-4-yl (wherein X⁷ and R²⁹ are as defined in claim 1);

15) 1-(R²⁹X⁸)but-2-yn-4-yl (wherein X⁸ and R²⁹ are as defined in claim 1);

16) C₂₋₃alkylX⁹C₁₋₃alkylR²⁹ (wherein X⁹ and R²⁹ are as defined in claim 1);

17) C₂₋₃alkylX⁹C₁₋₃alkylR²⁸ (wherein X⁹ and R²⁸ are as defined in claim 1);

18) C₂₋₅alkenyl which may be unsubstituted or which may be substituted with one or more fluorine atoms or with one or two groups selected from hydroxy, fluoro, amino, C₁₋₄alkylamino, N,N-di(C₁₋₄alkyl)amino, aminosulphonyl, N-C₁₋₄alkylaminosulphonyl and N,N-di(C₁₋₄alkyl)aminosulphonyl;

- 331 -

19) C_{2,3}alkynyl which may be unsubstituted or which may be substituted with one or more fluorine atoms or with one or two groups selected from hydroxy, fluoro, amino, C_{1,4}alkylamino, N,N-di(C_{1,4}alkyl)amino, aminosulphonyl, N-C_{1,4}alkylaminosulphonyl and N,N-di(C_{1,4}alkyl)aminosulphonyl;

5 20) C_{2,4}alkenylX⁹C_{1,3}alkylR²⁸ (wherein X⁹ and R²⁸ are as defined in claim 1);

21) C_{2,4}alkynylX⁹C_{1,3}alkylR²⁸ (wherein X⁹ and R²⁸ are as defined in claim 1); and

22) C_{1,3}alkylR⁵⁴(C_{1,3}alkyl)_q(X⁹)_rR⁵⁵ (wherein X⁹, q, r, R⁵⁴ and R⁵⁵ are as defined in claim 1);

and additionally wherein any C_{1,3}alkyl, C_{2,3}alkenyl or C_{2,3}alkynyl group in R⁵X¹ may bear one or more substituents selected from hydroxy, halogeno and amino].

10

11. A compound according to any one of claims 9 and 10 wherein Zb is -O-.

12. A compound according to any one of claims 9, 10 and 11 wherein ring C is a 9-10-membered heteroaromatic bicyclic moiety which contains 1-3 heteroatoms selected independently from O, N and S.

15

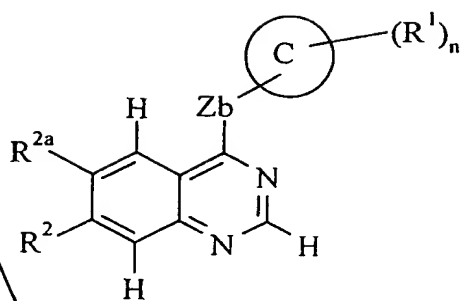
13. A compound according to any one of claims 9, 10, 11 and 12 wherein R¹ represents oxo, halogeno, hydroxy, C_{1,2}alkoxy, C_{1,2}alkyl, C_{1,2}alkoxymethyl, C_{2,3}alkanoyl, C_{1,2}haloalkyl, cyano, amino, C_{2,4}alkenyl, C_{2,4}alkynyl, C_{2,3}alkanoyloxy, nitro, C_{2,3}alkanoylamino, C_{1,2}alkoxycarbonyl, C_{1,2}alkylsulphanyl, C_{1,2}alkylsulphinyl, C_{1,2}alkylsulphonyl, carbamoyl, N-C_{1,2}alkylcarbamoyl, N,N-di(C_{1,2}alkyl)carbamoyl, aminosulphonyl, N-C_{1,2}alkylaminosulphonyl, N,N-di(C_{1,2}alkyl)aminosulphonyl, N-(C_{1,2}alkylsulphonyl)amino, N-(C_{1,2}alkylsulphonyl)-N-(C_{1,2}alkyl)amino or a C_{3,7}alkylene chain joined to two ring C carbon atoms.

25

14. A compound according to any one of claims 9, 10, 11, 12 and 13 wherein n is 0, 1 or 2.

15. A compound of the formula IIb:

30



(IIb)

15 [wherein:

ring C, R^1 , R^2 and n are as defined in claim 1, Zb is -O- and R^{2a} is as defined in claim 9 with the proviso that R^2 does not have any of the following values:

hydrogen, substituted or unsubstituted C_{1-3} alkyl, halogeno, C_{1-3} alkoxy, C_{2-5} alkenyl, phenoxy or phenyl C_{1-3} alkoxy;

20 or a salt thereof.

16. A compound according to claim 9 selected from

6-methoxy-7-((1-methylpiperidin-4-yl)methoxy)-4-(2-naphthyloxy)quinazoline,

6-methoxy-7-((1-methylpiperidin-4-yl)methoxy)-4-(quinolin-7-yloxy)quinazoline,

25 7-(3-(1,1-dioxothiomorpholino)propoxy)-6-methoxy-4-(quinolin-7-yloxy)quinazoline,

6-methoxy-7-(3-(4-methylpiperazin-1-yl)propoxy)-4-(quinolin-7-yloxy)quinazoline,

6-methoxy-7-((1-methylpiperidin-3-yl)methoxy)-4-(quinolin-7-yloxy)quinazoline,

4-(4-chloroquinolin-7-yloxy)-6-methoxy-7-(3-morpholinopropoxy)quinazoline,

6-methoxy-7-((1-methylpiperidin-4-yl)methoxy)-4-(4-methylquinolin-7-yloxy)quinazoline,

30 6-methoxy-4-(4-methylquinolin-7-yloxy)-7-(3-(pyrrolidin-1-yl)propoxy)quinazoline,

6-methoxy-7-(2-(2-methoxyethoxy)ethoxy)-4-(quinolin-7-yloxy)quinazoline,

- 6-methoxy-7-((1-(2-methylsulphonyl)ethyl)piperidin-4-yl)methoxy)-4-(quinolin-7-yloxy)quinazoline,
4-(2,3-dimethylindol-5-yloxy)-6-methoxy-7-(1-methylpiperidin-4-ylmethoxy)quinazoline,
4-(2,3-dimethylindol-5-yloxy)-6-methoxy-7-(3-pyrrolidin-1-ylpropoxy)quinazoline,
5 6-methoxy-7-(1-methylpiperidin-4-ylmethoxy)-4-(2-trifluoromethylindol-5-yloxy)quinazoline,
6-methoxy-7-(3-pyrrolidin-1-ylpropoxy)-4-(2-trifluoromethylindol-5-yloxy)quinazoline,
(*R,S*)-4-(3-fluoroquinolin-7-yloxy)-6-methoxy-7-((1-methylpiperidin-3-yl)methoxy)quinazoline,
10 4-(indol-5-yloxy)-6-methoxy-7-(3-methylsulphonylpropoxy)quinazoline,
7-(3-*N,N*-dimethylaminopropoxy)-6-methoxy-4-(2-methylindol-5-yloxy)quinazoline,
6-methoxy-4-(2-methylindol-5-yloxy)-7-(2-(2-morpholinoethoxy)ethoxy)quinazoline,
7-(2-(*N,N*-diethylamino)ethoxy)-6-methoxy-4-(2-methylindol-5-yloxy)quinazoline,
6-methoxy-7-(3-piperidinopropoxy)-4-(quinolin-7-yloxy)quinazoline,
15 4-(2-methylindol-5-yloxy)-7-(3-morpholinopropoxy)quinazoline,
4-(2-methylindol-5-yloxy)-7-(2-(piperidin-1-yl)ethoxy)quinazoline,
4-(2-methylindol-5-yloxy)-7-(2-(1*H*-1,2,4-triazol-1-yl)ethoxy)quinazoline,
6-methoxy-7-(3-piperidinopropoxy)-4-(6-trifluoromethylindol-5-yloxy)quinazoline,
7-(3-(methylsulphonyl)propoxy)-4-(2-methylindol-5-yloxy)quinazoline,
20 7-(3-(*N,N*-dimethylamino)propoxy)-4-(2,3-dimethylindol-5-yloxy)-6-methoxyquinazoline,
4-(2,3-dimethylindol-5-yloxy)-6-methoxy-7-(1-methylpiperidin-3-ylmethoxy)quinazoline,
7-(2-(*N,N*-diethylamino)ethoxy)-4-(indol-5-yloxy)-6-methoxyquinazoline,
4-(indol-5-yloxy)-6-methoxy-7-(2-(piperidin-2-yl)ethoxy)quinazoline,
4-(indol-5-yloxy)-6-methoxy-7-(2-(piperidin-1-yl)ethoxy)quinazoline,
25 4-(indol-6-yloxy)-6-methoxy-7-(3-morpholinopropoxy)quinazoline,
7-(3-(ethylsulphonyl)propoxy)-6-methoxy-4-(2-methylindol-5-yloxy)quinazoline,
6-methoxy-4-(3-methylindol-5-yloxy)-7-(3-piperidinopropoxy)quinazoline,
7-(2-hydroxy-3-piperidinopropoxy)-6-methoxy-4-(2-methylindol-5-yloxy)quinazoline,
7-(2-hydroxy-3-(4-methylpiperazin-1-yl)propoxy)-6-methoxy-4-(2-methylindol-5-yloxy)quinazoline,
30 6-methoxy-4-(2-methylindol-5-yloxy)-7-(2-(*N*-methylamino)ethoxy)quinazoline, and
7-(2-hydroxy-3-(isopropylamino)propoxy)-6-methoxy-4-(2-methylindol-5-yloxy)quinazoline,

or a salt thereof.

17. A compound according to claim 9 selected from

- 6-methoxy-7-(3-morpholinopropoxy)-4-(quinolin-7-yloxy)quinazoline,
5 6-methoxy-4-(2-methylindol-5-yloxy)-7-((1-methylpiperidin-4-yl)methoxy)quinazoline,
4-(indol-5-yloxy)-6-methoxy-7-(1-methylpiperidin-4-ylmethoxy)quinazoline,
4-(indol-5-yloxy)-6-methoxy-7-(3-pyrrolidin-1-ylpropoxy)quinazoline,
6-methoxy-4-(2-methylindol-5-yloxy)-7-(3-methylsulphonylpropoxy)quinazoline,
7-((1-cyanomethyl)piperidin-4-ylmethoxy)-6-methoxy-4-(2-methylindol-5-yloxy)quinazoline,
10 6-methoxy-4-(2-methylindol-5-yloxy)-7-(2-morpholinoethoxy)quinazoline,
6-methoxy-4-(2-methylindol-5-yloxy)-7-(2-pyrrolidin-1-ylethoxy)quinazoline,
6-methoxy-4-(2-methylindol-5-yloxy)-7-(1-methylpiperidin-3-ylmethoxy)quinazoline,
6-methoxy-4-(2-methylindol-5-yloxy)-7-(2-piperidinoethoxy)quinazoline,
6-methoxy-4-(2-methylindol-5-yloxy)-7-(2-(N-methyl-N-(4-
15 pyridyl)amino)ethoxy)quinazoline,
6-methoxy-4-(2-methylindol-5-yloxy)-7-(3-morpholinopropoxy)quinazoline,
6-methoxy-7-(2-(2-methoxyethoxy)ethoxy)-4-(2-methylindol-5-yloxy)quinazoline,
6-methoxy-4-(2-methylindol-5-yloxy)-7-(2-(1*H*-1,2,4-triazol-1-yl)ethoxy)quinazoline,
6-methoxy-4-(2-methylindol-5-yloxy)-7-(2-(2-(4-methylpiperazin-1-
20 yl)ethoxy)ethoxy)quinazoline,
6-methoxy-4-(2-methylindol-5-yloxy)-7-(3-piperidinopropoxy)quinazoline,
4-(indol-5-yloxy)-6-methoxy-7-(3-piperidinopropoxy)quinazoline
6-methoxy-7-(1-(2-methoxyethyl)piperidin-4-ylmethoxy)-4-(2-methylindol-5-
yloxy)quinazoline,
25 6-methoxy-4-(2-methylindol-5-yloxy)-7-((2-(2-pyrrolidin-1-
ylethyl)carbamoyl)vinyl)quinazoline,
6-methoxy-4-(2-methylindol-5-yloxy)-7-(3-(4-methylpiperazin-1-yl)propoxy)quinazoline,
6-methoxy-4-(2-methylindol-5-yloxy)-7-(piperidin-4-ylmethoxy)quinazoline,
6-methoxy-4-(2-methylindol-5-yloxy)-7-(2-(piperidin-4-yloxy)ethoxy)quinazoline,
30 6-methoxy-4-(2-methylindol-5-yloxy)-7-(2-(N-methyl-N-
methylsulphonylamino)ethoxy)quinazoline,

- 7-(2-(1-(2-cyanoethyl)piperidin-4-yloxy)ethoxy)-6-methoxy-4-(2-methylindol-5-yloxy)quinazoline,
4-(2-methylindol-5-yloxy)-7-(3-(pyrrolidin-yl)propoxy)quinazoline,
4-(2-methylindol-5-yloxy)-7-(3-(1,1-dioxothiomorpholino)propoxy)quinazoline,
5 4-(2-methylindol-5-yloxy)-7-(piperidin-4-ylmethoxy)quinazoline,
4-(indol-5-yloxy)-6-methoxy-7-(2-(2-methoxyethoxy)ethoxy)quinazoline,
7-(3-(N,N-dimethylamino)propoxy)-4-(indol-5-yloxy)-6-methoxyquinazoline,
7-(3-(N,N-diethylamino)propoxy)-4-(indol-5-yloxy)-6-methoxyquinazoline,
7-(3-(1,1-dioxothiomorpholino)propoxy)-4-(indol-5-yloxy)-6-methoxyquinazoline,
10 4-(indol-5-yloxy)-6-methoxy-7-(2-(4-pyridyloxy)ethoxy)quinazoline,
4-(indol-6-yloxy)-6-methoxy-7-(3-piperidinopropoxy)quinazoline,
7-(1-(2-methoxyethyl)piperidin-4-ylmethoxy)-4-(2-methylindol-5-yloxy)quinazoline,
7-(2-hydroxy-3-morpholinopropoxy)-6-methoxy-4-(2-methylindol-5-yloxy)quinazoline,
7-(2-(1-(2-methoxyethyl)piperidin-4-yl)ethoxy)-6-methoxy-4-(2-methylindol-5-yloxy)quinazoline,
15 7-(2-hydroxy-3-pyrrolidin-1-ylpropoxy)-6-methoxy-4-(2-methylindol-5-yloxy)quinazoline,
7-(3-(N,N-diethylamino)-2-hydroxypropoxy)-6-methoxy-4-(2-methylindol-5-yloxy)quinazoline,
7-(3-(1,1-dioxothiomorpholino)propoxy)-6-methoxy-4-(2-methylindol-5-yloxy)quinazoline,
20 6-methoxy-4-(2-methylindol-5-yloxy)-7-(2-(4-pyridyloxy)ethoxy)quinazoline,
4-(indol-5-yloxy)-6-methoxy-7-(3-morpholinopropoxy)quinazoline,
(2*R*)-6-methoxy-(2-methyl-1*H*-indol-5-yloxy)-7-(2-hydroxy-3-piperidinopropoxy)quinazoline,
(5*R*)-6-methoxy-4-(2-methyl-1*H*-indol-5-yloxy)-7-(2-oxopyrrolidin-5-ylmethoxy)quinazoline,
25 4-(4-bromoindol-5-yloxy)-6-methoxy-7-(3-piperidinopropoxy)quinazoline,
6-methoxy-4-(2-methylindol-5-yloxy)-7-(1-(2-(pyrrolidin-1-yl)ethyl)-piperidin-4-ylmethoxy)quinazoline,
(2*R*)-7-(2-hydroxy-3-(pyrrolidin-1-yl)propoxy)-4-(indol-5-yloxy)-6-methoxyquinazoline,
(2*R*)-7-(2-hydroxy-3-morpholinopropoxy)-4-(indol-5-yloxy)-6-methoxyquinazoline,
30 (2*R*)-7-(2-hydroxy-3-piperidinopropoxy)-4-(indol-5-yloxy)-6-methoxyquinazoline,
(2*S*)-7-(2-hydroxy-3-(N,N-diisopropyl)amino)propoxy)-4-(indol-5-yloxy)-6-methoxyquinazoline,

- (2*S*)-7-(2-hydroxy-3-piperidinopropoxy)-4-(indol-5-yloxy)-6-methoxyquinazoline,
(2*R*)-7-(2-hydroxy-3-piperidinopropoxy)-6-methoxy-4-(3-methylindol-5-yloxy)quinazoline,
(2*R*)-7-(2-hydroxy-3-(pyrrolidin-1-yl)propoxy)-6-methoxy-4-(3-methylindol-5-yloxy)quinazoline,
5 (2*R*)-7-(2-hydroxy-3-(pyrrolidin-1-yl)propoxy)-6-methoxy-4-(2-methylindol-5-yloxy)quinazoline,
(2*R*)-7-(2-hydroxy-3-(4-methylpiperazin-1-yl)propoxy)-6-methoxy-4-(2-methylindol-5-yloxy)quinazoline,
6-methoxy-4-(2-methylindol-5-yloxy)-7-(1-(2-morpholinoethyl)piperidin-4-ylmethoxy)quinazoline,
10 4-(3-fluoro-quinolin-7-yloxy)-6-methoxy-7-(3-piperidinopropoxy)quinazoline,
4-(3-fluoro-quinolin-7-yloxy)-6-methoxy-7-(3-(pyrrolidin-1-yl)propoxy)quinazoline,
6-methoxy-7-(3-(pyrrolidin-1-yl)propoxy)-4-(1*H*-pyrrolo[2,3-*b*]pyridin-5-yloxy)quinazoline,
(2*S*)-6-methoxy-(2-methyl-1*H*-indol-5-yloxy)-7-(2-hydroxy-3-piperidinopropoxy)quinazoline,
15 and
4-(6-fluoro-2-methylindol-5-yloxy)-6-methoxy-7-(3-(pyrrolidin-1-yl)propoxy)quinazoline,
or a salt thereof.

18. A compound according to claim 9 selected from
20 6-methoxy-4-(2-methylindol-5-yloxy)-7-(3-(pyrrolidin-1-yl)propoxy)quinazoline,
4-(4-fluoroindol-5-yloxy)-6-methoxy-7-(1-methylpiperidin-4-ylmethoxy)quinazoline,
4-(4-fluoroindol-5-yloxy)-6-methoxy-7-(3-(4-methylpiperazin-1-yl)propoxy)quinazoline,
4-(6-fluoroindol-5-yloxy)-6-methoxy-7-(3-(pyrrolidin-1-yl)propoxy)quinazoline,
4-(4-fluoroindol-5-yloxy)-6-methoxy-7-(3-(pyrrolidin-1-yl)propoxy)quinazoline,
25 4-(4-fluoroindol-5-yloxy)-6-methoxy-7-(3-piperidinopropoxy)quinazoline,
4-(4-fluoro-2-methylindol-5-yloxy)-6-methoxy-7-(3-(pyrrolidin-1-yl)propoxy)quinazoline,
4-(4-fluoro-2-methylindol-5-yloxy)-6-methoxy-7-(3-piperidinopropoxy)quinazoline,
4-(4-fluoro-2-methylindol-5-yloxy)-6-methoxy-7-((1-methylpiperidin-4-yl)methoxy)quinazoline,
30 4-(4-fluoro-2-methylindol-5-yloxy)-6-methoxy-7-(3-(4-methylpiperazin-1-yl)propoxy)quinazoline,
4-(4-fluoroindol-5-yloxy)-6-methoxy-7-(2-(1-methylpiperidin-4-yl)ethoxy)quinazoline,

- 337 -

(2*R*)-7-(2-hydroxy-3-(pyrrolidin-1-yl)propoxy)-4-(4-fluoro-2-methylindol-5-yloxy)-6-methoxyquinazoline, and

4-(4-fluoro-2-methylindol-5-yloxy)-6-methoxy-7-(2-(1-methylpiperidin-4-yl)ethoxy)quinazoline,

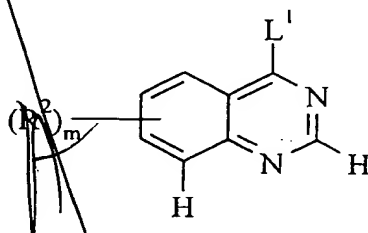
5 or a salt thereof.

19. A compound according to claim 9 in the form of a pharmaceutically acceptable salt.

10 20. A process for the preparation of a compound of formula I or salt thereof which comprises:

(a) the reaction of a compound of the formula III:

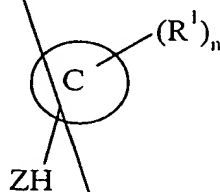
15



(III)

20

(wherein R^2 and m are as defined in claim 1 and L^1 is a displaceable moiety), with a compound of the formula IV:



(IV)

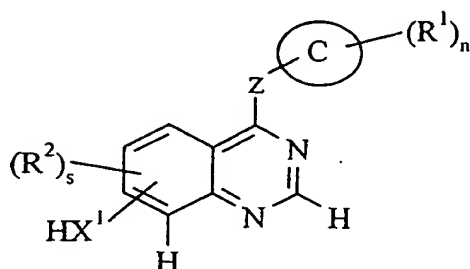
25

(wherein ring C, R^1 , Z and n are as defined in claim 1);

(b) a compound of formula I or a salt thereof wherein at least one R^2 is R^5X^1 wherein R^5 is as defined in claim 1 and X^1 is -O-, -S-, -OC(O)- or -NR¹⁰- (wherein R^{10} independently

- 338 -

represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl) may be prepared by the reaction of a compound of the formula V:



(V)

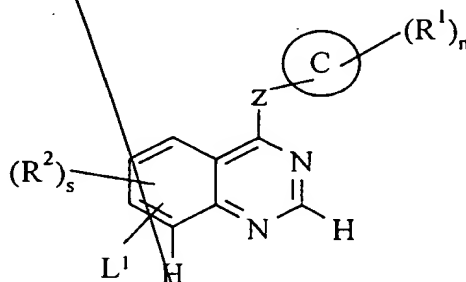
(wherein ring C, Z, R¹, R² and n are as defined in claim 1 and X¹ is as herein defined in this section and s is an integer from 0 to 2) with a compound of formula VI:



(VI)

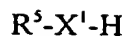
(wherein R⁵ is as defined in claim 1 and L¹ is as herein defined);

(c) a compound of the formula I or a salt thereof wherein at least one R² is R⁵X¹ wherein R⁵ is as defined in claim 1 and X¹ is -O-, -S-, -OC(O)- or -NR¹⁰- (wherein R¹⁰ represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl) may be prepared by the reaction of a compound of the formula VII:



(VII)

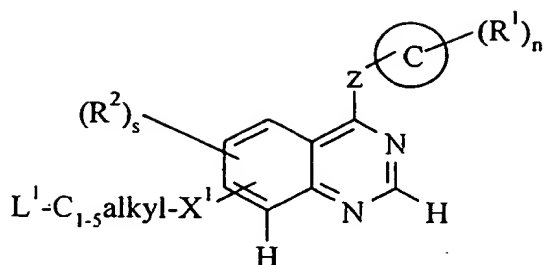
with a compound of the formula VIII:



(VIII)

(wherein R^1 , R^2 , R^5 , ring C, Z and n are as defined in claim 1 and L^1 , s and X^1 are as herein defined);

- 5 (d) a compound of the formula I or a salt thereof wherein at least one R^2 is R^5X^1 wherein X^1 is as defined in claim 1 and R^5 is $C_{1-5}alkylR^{113}$, wherein R^{113} is selected from one of the following nine groups:
- 1) $X^{19}C_{1-3}alkyl$ (wherein X^{19} represents -O-, -S-, -SO₂-, -NR¹¹⁴C(O)- or -NR¹¹⁵SO₂- (wherein R^{114} and R^{115} which may be the same or different are each hydrogen, $C_{1-3}alkyl$ or $C_{1-3}alkoxyC_{2-3}alkyl$);
 - 10 2) NR¹¹⁶R¹¹⁷ (wherein R^{116} and R^{117} which may be the same or different are each hydrogen, $C_{1-3}alkyl$ or $C_{1-3}alkoxyC_{2-3}alkyl$);
 - 3) $X^{20}C_{1-5}alkylX^5R^{22}$ (wherein X^{20} represents -O-, -S-, -SO₂-, -NR¹¹⁸C(O)-, -NR¹¹⁹SO₂- or -NR¹²⁰- (wherein R^{118} , R^{119} , and R^{120} which may be the same or different are each hydrogen, $C_{1-3}alkyl$ or $C_{1-3}alkoxyC_{2-3}alkyl$) and X^5 and R^{22} are as defined in claim 1);
 - 15 4) R^{28} (wherein R^{28} is as defined in claim 1);
 - 5) $X^{21}R^{29}$ (wherein X^{21} represents -O-, -S-, -SO₂-, -NR¹²¹C(O)-, -NR¹²²SO₂-, or -NR¹²³- (wherein R^{121} , R^{122} , and R^{123} which may be the same or different are each hydrogen, $C_{1-3}alkyl$ or $C_{1-3}alkoxyC_{2-3}alkyl$) and R^{29} is as defined in claim 1); and
 - 20 6) $X^{22}C_{1-3}alkylR^{29}$ (wherein X^{22} represents -O-, -S-, -SO₂-, -NR¹²⁴C(O)-, -NR¹²⁵SO₂- or -NR¹²⁶- (wherein R^{124} , R^{125} and R^{126} each independently represents hydrogen, $C_{1-3}alkyl$ or $C_{1-3}alkoxyC_{2-3}alkyl$) and R^{29} is as defined in claim 1);
 - 7) R^{29} (wherein R^{29} is as defined in claim 1);
 - 8) $X^{22}C_{1-4}alkylR^{28}$ (wherein X^{22} and R^{28} are as defined in claim 1); and
 - 25 9) $R^{54}(C_{1-4}alkyl)_q(X^9)_rR^{55}$ (wherein q, r, X^9 , R^{54} and R^{55} are as defined in claim 1);
- may be prepared by reacting a compound of the formula IX:



(IX)

- 10 (wherein X^1 , R^1 , R^2 , ring C, Z and n are as defined in claim 1 and L^1 and s are as herein defined) with a compound of the formula X:



(X)

(wherein R^{113} is as defined herein);

- 15 (e) a compound of the formula I or a salt thereof wherein one or more of the substituents $(R^2)_m$ is represented by $-NR^{127}R^{128}$, where one (and the other is hydrogen) or both of R^{127} and R^{128} are C_{1-3} alkyl, may be effected by the reaction of compounds of formula I wherein the substituent $(R^2)_m$ is an amino group and an alkylating agent; or
- (f) a compound of the formula I or a salt thereof wherein X^1 is $-SO-$ or $-SO_2-$ may be
- 20 prepared by oxidation from the corresponding compound in which X^1 is $-S-$ or $-SO-$; and when a salt of a compound of formula I is required, reaction of the compound obtained with an acid or base whereby to obtain the desired salt.

21. A pharmaceutical composition which comprises as active ingredient a compound
- 25 of formula I or a pharmaceutically acceptable salt thereof according to claim 9 in association with a pharmaceutically acceptable excipient or carrier.

22. A method for producing an antiangiogenic and/or vascular permeability reducing effect in a warm-blooded animal in need of such treatment which comprises administering to
- 30 said animal an effective amount of a compound of formula I as defined in claim 1 or a pharmaceutically acceptable salt thereof.

PAT 34 AMBT

23. A compound 4-fluoro-5-hydroxy-2-methylindole or a salt thereof.
24. A compound 4-fluoro-5-hydroxyindole or a salt thereof.
- 5 25. A compound 6-fluoro-5-hydroxy-2-methylindole or a salt thereof.
26. A compound 6-fluoro-5-hydroxyindole or a salt thereof.
- 10 27. A process for the preparation of 4-fluoro-5-hydroxy-2-methylindole according to any one of those described in Example 237.
28. A process for the preparation of 4-fluoro-5-hydroxyindole as described in Example 242.
- 15 29. A process for the preparation of 6-fluoro-5-hydroxyindole as described in Example 242.
- 20 30. A process for the preparation of 6-fluoro-5-hydroxy-2-methylindole as described in Example 250.

add
A22

AMENDED SHEET